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Original Article

Sleep duration and growth outcomes across the first two years of life in the GUSTO study

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ABSTRACT

Background and Aim: Short sleep duration is thought to be a factor contributing to increased body mass index (BMI) in both school-age children and adults. Our aim was to determine whether sleep duration associates with growth outcomes during the first two years of life.

Study design: Participants included 899 children enrolled in the Growing Up in Singapore Towards healthy Outcomes (GUSTO) birth cohort study. Anthropometric data (weight and body length) and parental reports of sleep duration were collected at 3, 6, 9, 12, 18, and 24 months of age. A mixed-model analysis was used to evaluate the longitudinal association of BMI and body length with sleep duration. In subgroup analyses, effects of ethnicity (Chinese, Indian, and Malay) and short sleep at three months of age (≤12 h per day) were examined on subsequent growth measures.

Results: In the overall cohort, sleep duration was significantly associated with body length ($\beta = 0.028$, 95% confidence interval [CI] 0.002–0.053, $p = 0.033$), but not BMI, after adjustment for potential confounding factors. Only in Malay children, shorter sleep was associated with a higher BMI ($\beta = -0.042$, 95% CI −0.071 to −0.012, $p = 0.005$) and shorter body length ($\beta = 0.079$, 95% CI 0.030–0.128, $p = 0.002$). In addition, shorter sleep was associated with a higher BMI and shorter body length in children who slept ≤12 h per day at three months of age.

Conclusion: The association between sleep duration and growth outcomes begins in infancy. The small but significant relationship between sleep and growth anthropometric measures in early life might be amplified in later childhood.

Abbreviations: BISQ, Brief Infant Sleep Questionnaire; BMI, body mass index; GUSTO, Growing Up in Singapore Towards healthy Outcomes.

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1. Introduction

Over the past few decades, the rate of childhood obesity has increased substantially. Today, about one-third of children worldwide are either overweight or obese [1]. In parallel with the global obesity epidemic, there has been a gradual decline in sleep duration over the past several decades that includes children in Asia, Europe, and the United States [2]. Numerous studies have demonstrated an inverse correlation between sleep duration and adiposity or obesity risk in children and adults [3–10]. Possible mechanisms include decreased leptin and increased ghrelin associated with sleep deprivation [11], leading to increased caloric intake and reduced energy expenditure, both of which contribute to obesity. In addition, sleep is thought to be essential for growth during child development. As pulsatile release of human growth hormone occurs during slow-wave sleep [12], it is possible that chronic exposure to short sleep during childhood affects growth (body length), although this hypothesis has yet to be systematically tested.

The relationship between sleep duration and growth outcomes has been primarily investigated in school-age children and not in younger age groups [3,6,7,9,10,13]. In addition, earlier studies were performed in populations that are predominantly Caucasian, even though epidemiologic evidence indicates that sleep duration in children is shorter in East Asian countries [14–16]. Similarly, in Singapore it has been reported that 2-year-old children sleep about 2 h less per day relative to Swiss children of the same age [17,18], but the impact on growth outcomes has not been explored. This study, therefore investigated the relationship between sleep duration and growth in Singaporean children during the first two years of life. Here, we tested the hypothesis that shorter sleep duration is associated with a higher body mass index (BMI) and shorter body length across early development.

2. Methods

2.1. Study design and population

This study was conducted as part of the Growing Up in Singapore Towards healthy Outcomes (GUSTO) birth cohort study, which aims to identify perinatal and early-life factors that affect growth and metabolic health of children raised in Singapore. The GUSTO study methodology has been described in detail elsewhere [19]. Pregnant women aged ≥18 years were recruited in their first trimester from KK Women's and Children's Hospital and the National University Hospital, the two major public hospitals with obstetric services in Singapore. Only women with a homogeneous parental ethnic background who were Chinese, Malay, or Indian were eligible for the study. Women were excluded if they were on chemotherapy, taking psychotropic drugs, or if they had preexisting diabetes mellitus. A total of 1247 women with singleton pregnancy were recruited, with children born between 30 November 2009 and 1 May 2011. Informed written consent was obtained from each participant on the day of study enrollment, and procedures were approved by the National Healthcare Group Institutional Review Board (IRB) and the SingHealth Centralized IRB. This study is registered under the Clinical Trials Identifier NCT01174875.

2.2. Sleep duration

The Brief Infant Sleep Questionnaire (BISQ) was used to assess infant sleep behavior at 3, 6, 9, 12, 18, and 24 months of age [20]. The BISQ was administered to the main caregiver of the child (mostly mothers) in English, Chinese, Tamil, or Malay language. Two items were selected from the questionnaire responses: (1) "How much time (on average) does your child spend in sleep during the NIGHT?" and (2) "How much time (on average) does your child spend in sleep (naps) during the DAY?" Both responses were reported in hours and minutes. The main exposure parameter in this study was total daily sleep duration, which was calculated as the sum of daytime sleep and nighttime sleep.

2.3. Growth measures

Growth measures were examined at time points corresponding to administration of the BISQ. Weight was measured to the nearest gram using a calibrated scale (SECA 334 weighing scale; SECA Corp., Hamburg, Germany). Recumbent body length was measured in duplicate to the nearest 0.1 cm from the top of the head to the soles of the feet using a measuring mat (SECA 210 mobile measuring mat; SECA Corp.). BMI was calculated as the weight in kilograms divided by squared body length in meters.

2.4. Covariates

Demographic parameters such as ethnicity, maternal education, and household income were collected at the beginning of the study. In addition, parental anthropometric measures were collected during pregnancy, and maternal height and BMI at 26 weeks of gestation were used in this analysis. Data concerning delivery and perinatal risk factors included sex of the child, gestational age, birth weight and length, pregnancy smoking status, and maternal gestational diabetes mellitus. Anthropometric measurements of newborns were completed within 24 h of birth. Breast-feeding status was documented at every visit during the study as exclusive, predominant, partial, or formula only, which followed the World Health Organization's definitions [21]. Considering that breast-feeding patterns changed over time, a weighted sum was used (weights: exclusive breast-feeding = 1, predominant breast-feeding = 0.75, partial breast-feeding = 0.5, and formula only = 0) for total breast-feeding duration which took into account the type of breast-feeding and its corresponding duration. Early lifestyle measures in children were also included as covariates such as total media use and outdoor physical activity at 24 months. Data on the number of daily hours spent on the computer and handheld devices or watching television were collected at 24 months of age for both weekdays and weekends, and a weighted average of total media use was calculated. The average number of hours per day spent playing/exercising outdoors was similarly calculated after taking into account weekday and weekend behavior.

2.5. Statistical analysis

We first examined maternal and offspring characteristics, as well as exposure and outcome variables, of the three main ethnic groups in Singapore (Chinese, Indian, and Malay). Mean comparisons among groups were performed using one-way analysis of variance (ANOVA). In order to account for the dependence among repeated measures across time points, as well as to maximize usage of longitudinal data, a mixed model without random effects was used to examine the relationship between BMI and sleep duration from three months to two years of age. Sleep duration and BMI values at 3, 6, 9, 12, 18, and 24 months were entered into the mixed model, together with covariates described in the previous section. A similar model was used to examine the longitudinal relationship between body length and sleep duration.

In subgroup analyses, the associations of BMI and body length with sleep duration within different ethnic groups were examined after adjusting for covariates. Similarly, children within subgroups with either shorter or longer sleep duration were also examined, with shorter sleep duration defined as ≤12 h per day at three months of age. This cutoff was chosen because it was the mean sleep duration at the 3-month time point. In addition, our preliminary analyses suggested stronger ethnic differences in sleep behavior and more variable sleep duration between children at 3 months of age, as compared to later time points. Therefore, the study focused on determining whether sleep duration reported at 3 months was associated with growth outcomes in later development. Data were analyzed using IBM SPSS Statistics for Windows, V. 19.0 (IBM Corp; Armonk, NY, USA).

3. Results

Of the 1247 mother–infant pairs recruited at the beginning of the study, 348 subjects were excluded due to missing BISQ or growth data. The remaining 899 subjects had at least one time point with outcome (BMI and body length) and exposure (sleep duration) measures. This group consisted of a greater proportion of mothers with Chinese ethnicity, higher education, and higher household income in comparison to those who were excluded based on missing data (Supplementary Table S1). For participants with available sleep and growth data, maternal and offspring characteristics differed substantially across ethnic groups (Table 1). As compared to Chinese and Indian mothers, Malay women were of lower socioeconomic status in terms of education and income level, and they had a higher BMI at 26 weeks of gestational age. In addition, the duration of breast-feeding was the lowest and media use at 24 months of age was the highest in the Malay group. Across nearly all time points examined, Malay children exhibited shorter body length and higher BMI values in comparison to children in the other ethnic groups (Table 2).

On average, sleep duration decreased by about an hour (12.08–11.14 h) from three months to 24 months of age (Table 3). At three months, sleep duration was about 1.5 h shorter in Malay children in comparison to Chinese and Indian children. At 6 months, sleep duration was about an hour longer in Chinese infants in comparison to the other groups, but these ethnic differences were not present at later time points. Based on the mixed-model analysis, sleep duration was not associated with BMI for the first two years of life, independent of adjustment for covariates (Table 4). By contrast, although the effect size was very small (Table 4), sleep duration was

Table 1
Maternal and offspring characteristics. Data are shown as mean (SD) or %.

	Chinese ($n = 507$)	Indian ($n = 155$)	Malay ($n = 237$)	Total ($n = 899$)	p-value
Mothers					
Highest education obtained (%)					
Below “A” level/diploma	31.01%	27.03%	68.58%	40.19%	<0.001
Household income (%)					
Below S$4000 per month	32.27%	50.35%	70.31%	45.67%	<0.001
Anthropometrics at 26 weeks gestation					
Weight (kg)	62.95 (9.61)	67.99 (11.60)	69.40 (14.13)	65.51 (11.67)	<0.001
Height (cm)	158.93 (5.67)	157.73 (5.44)	157.15 (5.71)	158.26 (5.69)	<0.001
BMI (kg/m^2)	24.68 (4.04)	26.97 (5.45)	27.81 (5.81)	25.90 (5.01)	<0.001
Perinatal risk factors					
Smoking during pregnancy (%)	1.79%	1.31%	5.11%	2.58%	0.017
Gestational diabetes (%)	19.13%	20.00%	10.13%	16.91%	0.005
Intrauterine growth retardation (%)	1.97%	5.16%	1.27%	2.34%	0.032
Offspring					
Birth information					
Male (%)	53.45%	48.39%	54.43%	52.84%	0.462
Gestational age (weeks)	38.82 (1.33)	38.71 (1.62)	38.52 (1.26)	38.72 (1.37)	0.023
Birth weight (kg)	3.13 (0.43)	3.05 (0.51)	3.13 (0.43)	3.12 (0.45)	0.126
Birth length (cm)	48.87 (2.27)	48.82 (2.19)	48.38 (2.07)	48.73 (2.21)	0.014
Birth BMI (kg/m^2)	13.08 (1.29)	12.74 (1.52)	13.31 (1.28)	13.08 (1.34)	<0.001
NICU stay (%)	2.76%	2.58%	3.38%	2.89%	0.869
Lifestyle risk factors					
Breast-feeding duration (months)	3.51 (3.49)	3.18 (3.27)	1.92 (2.71)	3.03 (3.33)	<0.001
Total media use at 24 months (h/day)	1.81 (1.89)	2.11 (2.24)	2.77 (2.58)	2.11 (2.19)	<0.001
Physical activity at 24 months (h/day)	0.80 (0.78)	0.92 (0.79)	0.73 (0.78)	0.80 (0.78)	0.053

Table 2
Growth outcomes in children. Data are shown as mean (SD).

Age and growth outcomes		Chinese ($n = 507$)	Indian ($n = 155$)	Malay ($n = 237$)	Total ($n = 899$)	p-value
3 months	Weight (kg)	6.31 (0.78)	5.82 (0.68)	6.01 (0.77)	6.15 (0.79)	<0.001
	Length (cm)	61.45 (2.49)	60.91 (2.20)	60.01 (2.34)	60.98 (2.48)	<0.001
	BMI (kg/m^2)	16.69 (1.50)	15.65 (1.42)	16.66 (1.58)	16.51 (1.55)	<0.001
6 months	Weight (kg)	7.81 (0.94)	7.53 (0.87)	7.67 (0.97)	7.73 (0.94)	0.005
	Length (cm)	67.43 (2.72)	67.39 (2.54)	66.15 (2.66)	67.10 (2.73)	<0.001
	BMI (kg/m^2)	17.15 (1.56)	16.55 (1.50)	17.53 (1.84)	17.14 (1.65)	<0.001
9 months	Weight (kg)	8.64 (1.00)	8.57 (0.99)	8.56 (1.03)	8.61 (1.01)	0.550
	Length (cm)	71.98 (2.92)	71.99 (2.66)	70.64 (2.89)	71.64 (2.93)	<0.001
	BMI (kg/m^2)	16.65 (1.45)	16.51 (1.45)	17.12 (1.54)	16.75 (1.49)	<0.001
12 months	Weight (kg)	9.38 (1.05)	9.46 (1.20)	9.29 (1.12)	9.37 (1.09)	0.354
	Length (cm)	75.78 (3.10)	76.30 (2.89)	74.22 (2.94)	75.48 (3.12)	<0.001
	BMI (kg/m^2)	16.31 (1.31)	16.20 (1.40)	16.81 (1.50)	16.42 (1.40)	<0.001
18 months	Weight (kg)	10.69 (1.24)	11.01 (1.52)	10.64 (1.38)	10.73 (1.33)	0.025
	Length (cm)	82.24 (3.33)	83.29 (3.45)	81.14 (3.21)	82.10 (3.39)	<0.001
	BMI (kg/m^2)	15.78 (1.50)	15.89 (1.55)	16.02 (1.46)	15.86 (1.50)	0.175
24 months	Weight (kg)	11.94 (1.50)	12.16 (1.72)	11.92 (1.60)	11.97 (1.57)	0.318
	Length (cm)	88.00 (3.66)	88.47 (3.68)	86.39 (3.21)	87.64 (3.63)	<0.001
	BMI (kg/m^2)	15.39 (1.32)	15.39 (1.52)	15.86 (1.48)	15.52 (1.41)	0.001

Table 3
Total daily sleep duration in hours during the first two years of life. Data are shown as mean (SD).

Age	Chinese	Indian	Malay	Total	*p*-value
3 months ($n = 643$)	12.49 (3.84)	12.43 (4.36)	10.96 (3.55)	12.08 (3.91)	<0.001
6 months ($n = 719$)	12.17 (2.65)	11.22 (2.95)	11.12 (3.08)	11.75 (2.85)	<0.001
9 months ($n = 547$)	11.78 (2.38)	11.74 (2.49)	11.30 (2.84)	11.67 (2.51)	0.170
12 months ($n = 515$)	11.69 (1.87)	12.09 (2.54)	11.32 (2.48)	11.66 (2.13)	0.047
18 months ($n = 411$)	11.60 (1.95)	11.47 (1.76)	11.58 (2.12)	11.57 (1.97)	0.917
24 months ($n = 399$)	11.23 (2.01)	11.19 (2.06)	10.92 (2.18)	11.14 (2.06)	0.414

Table 4
Longitudinal mixed-model analysis for BMI and body length versus sleep duration between 3 and 24 months of age, with adjustment for covariates. Values show estimates of fixed effects with 95% confidence intervals. $^{*}p < 0.05$ and $^{**}p < 0.01$.

	BMI (kg/m^2) ($n = 799$[a])		Body length (cm) ($n = 797$[a])	
	Univariate model	Multivariate model	Univariate model	Multivariate model
Sleep duration (h/day)	–0.009 (–0.026, 0.007)	–0.014(–0.030, 0.001)	0.039 (0.006, 0.071)*	0.028 (0.002, 0.053)*
Sex (male)	0.520 (0.340, 0.701)**	0.399 (0.227, 0.570)**	1.814 (1.462, 2.166)**	1.437 (1.155, 1.720)**
Ethnicity (Chinese)	–0.373 (–0.585, –0.162)**	–0.250 (–0.475, –0.025)*	1.438 (1.008, 1.868)**	1.122 (0.762, 1.482)**
Ethnicity (Indian)	–0.660 (–0.944, –0.375)**	–0.622 (–0.900, –0.345)**	1.354 (0.779, 1.929)**	1.085 (0.635, 1.535)**
Maternal education (Below "A" level/diploma)	–0.007 (–0.194, 0.181)	–0.107 (–0.320, 0.105)	–1.003 (–1.383, –0.624)**	–0.370 (–0.720, –0.020)*
Household income (Below S$4000/month)	–0.072 (–0.256, 0.113)	–0.055 (–0.259, 0.149)	–0.971 (–1.342, –0.600)**	0.003 (–0.340, 0.334)
Pregnancy smoking (no)	–0.288 (–0.852, 0.275)	–0.444 (–0.966, 0.079)	0.887 (–0.344, 2.118)	–0.091 (0.969, 0.788)
Gestational diabetes (no)	0.079 (–0.170, 0.328)	0.077 (–0.156, 0.309)	0.111 (–0.403, 0.625)	0.151 (–0.229, 0.532)
Birth weight (kg)	0.889 (0.689, 1.089)**	0.929 (0.692, 1.167)**	NA[b]	NA[b]
Birth length (cm)	NA[b]	NA[b]	0.515 (0.439, 0.590)**	0.427 (0.353, 0.501)**
Gestational age (weeks)	0.008 (–0.063, 0.079)	–0.142 (–0.218, –0.065)**	0.250 (0.105, 0.394)**	–0.164 (–0.287, –0.042)**
Maternal BMI (kg/m^2)	0.037 (0.019, 0.055)**	0.017 (–0.001, 0.036)	NA[b]	NA[b]
Maternal height (cm)	NA[b]	NA[b]	0.134 (0.102, 0.166)**	0.091 (0.065, 0.116)**
Breast-feeding duration (months)	0.004 (–0.023, 0.032)	0.018 (–0.009, 0.046)	0.016 (–0.040, 0.071)	–0.059 (–0.104, –0.013)*
Total media use at 24 months (h/day)	0.057 (0.015, 0.098)**	0.042 (0.002, 0.082)*	–0.028 (–0.116, 0.060)	0.024 (–0.044, 0.091)
Physical activity at 24 months (h/day)	0.056 (–0.061, 0.173)	0.024 (–0.088, 0.136)	0.220 (–0.022, 0.462)	0.047 (–0.139, 0.233)
Age 3 months[c]	0.925 (0.774, 1.076)**	0.953 (0.802, 1.104)**	–26.69 (–26.97, –26.41)**	–26.69 (–26.97, –26.40)**
Age 6 months[c]	1.626 (1.484 1.767)**	1.646 (1.505, 1.787)**	–26.65 (–20.91, –20.38)**	–20.64 (–20.91, –20.38)**
Age 9 months[c]	1.185 (1.043, 1.327)**	1.200 (1.059, 1.342)**	–16.04 (–16.30, –15.77)**	–16.02 (–16.29, –15.75)**
Age 12 months[c]	0.813 (0.684, 0.942)**	0.826 (0.697, 0.955)**	–12.11 (–12.37, –11.85)**	–12.11 (–12.37, –11.84)**
Age 18 months[c]	0.212 (0.094, 0.331)**	0.218 (0.099, 0.338)**	–5.39 (–5.62, –5.15)**	–5.37 (–5.61, –5.13)**

[a] The sample size is smaller than the starting sample ($n = 899$) due to the built-in modeling mechanism of the mixed model to exclude subjects with incomplete/insufficient longitudinal data for the model to give an accurate estimation of fixed effects.
[b] The respective variable is not applicable to the analysis, since only birth weight and maternal BMI are relevant for infant BMI, and only birth length and maternal height are relevant for infant body length.
[c] The reference is 24 months of age.

positively associated with body length for the first two years of life in both univariate ($\beta = 0.039$, 95% confidence interval (CI) 0.006–0.071 cm/h) and multivariate models ($\beta = 0.028$, 95% CI 0.002–0.053 cm/h). Other factors that associated positively with body length during early development included male sex, Chinese or Indian ethnicity, and greater birth length and maternal height.

In subgroup analyses based on ethnicity, sleep duration was negatively associated with BMI only in Malay children with each additional hour of sleep corresponding to a 0.042 kg/m^2 (95% CI –0.071 to –0.012, $p = 0.005$) decrease in BMI (Table 5). In addition, sleep duration was positively associated with body length in Malay subjects ($\beta = 0.079$, 95% CI 0.030–0.128 cm/h, $p = 0.002$). As average sleep duration was significantly less in Malay children in the first few months after delivery (Table 5), the possibility that the relationship between sleep duration and growth outcomes might be related to the greater number of shorter sleepers in the Malay group was considered. To investigate this, a subgroup analysis by sleep duration was performed, in which children were categorized as shorter sleepers (≤12 h) or longer sleepers (>12 h) based on parent-reported sleep behavior at 3 months of age (Table 5). Based on this definition, approximately two-thirds of Malay children were shorter sleepers (67.7%), whereas fewer than half of Chinese and Indian infants had shorter sleep duration (46.1% and 45.7%, respectively). In the shorter sleeper subgroup, sleep duration was negatively associated with BMI ($\beta = -0.036$, 95% CI –0.065 to –0.008 kg/m^2/h, $p = 0.013$) and positively associated with body length ($\beta = 0.070$, 95% CI 0.025–0.116 cm/h, $p = 0.003$), after adjustment for covariates. When analyzed further by ethnic subgroups, these relationships for BMI and body length remained significant in Chinese and Malay shorter sleepers (Table 5). In contrast to the shorter sleepers, the same analyses repeated on subgroups with longer-duration sleep did not show a significant association between sleep duration and BMI or body length.

Table 5
Longitudinal mixed model analysis of BMI and body length versus sleep duration, assessed in different ethnic subgroups and by sleep duration at 3 months of age, after adjustment for covariates. $^{*}p < 0.05$ and $^{**}p < 0.01$.

Subgroup and measures		Chinese	Indian	Malay	Overall
By ethnic group, BMI vs. sleep duration	β	–0.006	0.002	–0.042**	–0.014
	n	445	137	217	799
By ethnic group, body length vs. sleep duration	β	–0.006	–0.011	0.079**	0.028*
	n	445	137	216	797
Shorter sleepers (≤12 h/day), BMI vs. sleep duration	β	–0.044*	–0.061	–0.048*	–0.036*
	N	150	44	104	298
Shorter sleepers (≤12 h/day), body length vs. sleep duration	β	0.062*	0.108	0.081*	0.070**
	N	150	44	104	298
Longer sleepers (>12 h/day), BMI vs. sleep duration	β	0.008	–0.036	–0.043	–0.009
	n	175	50	49	274
Longer sleepers (>12 h/day), body length vs. sleep duration	β	0.024	–0.001	–0.034	0.006
	n	174	50	49	273

4. Discussion

Previous studies conducted in older children showed that sleep duration was negatively associated with BMI and body fat [3–10]. Sleep duration in childhood could have long-term consequences, as shorter sleep duration in children aged 5–11 years was associated with higher BMI values during adulthood in a prospective birth cohort study ($\beta = -0.99$, assessed at 32 years of age) [5]. In addition, based on a recent meta-analysis of 11 pediatric studies, a pooled odds ratio of 1.89 was observed for being overweight in children with short-duration sleep [3]. These studies suggest that sleep duration is an important contributing factor to metabolic health, which is also supported by studies in adult populations [11]. In this study, sleep duration did not associate with BMI in the full sample, which could be attributed to the much younger age of the participants. By comparison, a weak but significant association between sleep duration and BMI was observed in Malay participants, after controlling for differences in demographic and lifestyle factors. It is possible that the higher proportion of shorter sleepers in the Malay subgroup accounted for this observation given that both Chinese and Malay children who slept ≤12 h at 3 months of age showed a significant negative association between sleep duration and BMI. For children with sleep durations exceeding 12 h, there was no association with BMI, suggesting a possible ceiling effect when sufficient sleep is achieved.

In this study, sleep duration associated positively with body length during the first 2 years of life after adjusting for factors known to influence growth outcomes. Previous studies on body height have focused primarily on the relationship between sleep-disordered breathing and growth in children aged 2–6 years with adenotonsillar hypertrophy [22]. Following adenotonsillectomy, there is an improvement in sleep and an increase in height. The effects of sleep-disordered breathing on growth are considered to be mediated, at least in part, through disruption of slow-wave sleep when growth hormone is preferentially secreted [12]. While it might be predicted that shorter-duration sleep in healthy children would associate with shorter height, a weak negative association was observed between sleep duration and height in English and Scottish children aged 5–11 years whose sleep behavior was assessed by questionnaires [23]. Additional studies are therefore needed to establish whether sleep duration affects the rate of growth across childhood.

Although this study included several strengths, including enrollment of subjects across three major ethnic groups in Asia, multiple measures of sleep and growth outcomes over the first 2 years of development, and adjustment for known confounders, there were also limitations that should be considered. First, the majority of children (84.2%) did not have sleep duration data at all six time points. Thus, while the sampling frequency of sleep behavior was higher in comparison to other birth cohort studies, the response rate was also lower, and the sample size was relatively small at each time point (3234 responses were collected over six time points). The amount of missing data precluded accurate determination of growth trajectories and growth velocities on a per-individual basis. Therefore, this study did not examine whether the rate of growth varied with changes in sleep duration across the first 2 years of life. Another limitation of this study is that data on growth percentiles was not available for the three major ethnic groups examined in Singapore. As the sample sizes for the ethnic groups studied were too small to generate robust longitudinal growth percentiles, it was not possible to normalize growth parameters by ethnicity. The effect size, as measured by the β-coefficient, was also relatively small for sleep duration on growth outcomes, in comparison to other factors such as sex and ethnicity. Finally, it needs to be highlighted that sleep duration data collected by questionnaires are prone to parental estimation errors, and more reliable estimates of sleep behavior could potentially be obtained using objective measures, for example, actigraphy or polysomnography.

5. Conclusion

This study demonstrated that sleep duration associates weakly with anthropometric measures in early life. During the first 2 years of development, shorter sleep duration associated with higher BMI values in Malay participants and in the subgroup of children who slept ≤12 h per day at 3 months of age. In addition, shorter sleep was associated with shorter body length. Building on previous studies conducted in older children, these results suggest that sleep duration might begin to influence body weight and length from infancy. Future work should examine the impact of sleep duration on growth trajectories and velocities as our cohort matures longitudinally.

Conflict of interest

PDG, CYS, and KMG have received reimbursement for speaking at conferences sponsored by companies selling nutritional products, and they are part of an academic consortium that has received research funding from Abbott Nutrition, Nestle, and Danone. The remaining authors have no conflicts of interest or financial relationships to disclose.

The ICMJE Uniform Disclosure Form for Potential Conflicts of Interest associated with this article can be viewed by clicking on the following link: http://dx.doi.org/10.1016/j.sleep.2015.07.006.

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Appendix: Supplementary material

Supplementary data to this article can be found online at doi:10.1016/j.sleep.2015.07.006.

References

[1] Bethell C, Simpson L, Stumbo S, et al. National, state, and local disparities in childhood obesity. Health Aff 2010;29:347–56.
[2] Matricciani L, Olds T, Petkov J. In search of lost sleep: secular trends in the sleep time of school-aged children and adolescents. Sleep Med Rev 2012;16:203–11.
[3] Cappuccio FP, Taggart FM, Kandala N-B, et al. Meta-Analysis of Short Sleep Duration and Obesity in Children and Adults. Sleep 2008;31:619–26.
[4] Carter PJ, Taylor BJ, Williams SM, et al. Longitudinal analysis of sleep in relation to BMI and body fat in children: the FLAME study. Br Med J 2011;342:d2712.
[5] Landhuis CE, Poulton R, Welch D, et al. Childhood sleep time and long-term risk for obesity: a 32-year prospective birth cohort study. Pediatrics 2008;122:955–60.
[6] Magee L, Hale L. Longitudinal associations between sleep duration and subsequent weight gain: a systematic review. Sleep Med Rev 2012;16:231–41.

[7] Marshall NS, Glozier N, Grunstein RR. Is sleep duration related to obesity? A critical review of the epidemiological evidence. Sleep Med Rev 2008;12:289–98.
[8] Reilly JJ, Armstrong J, Dorosty AR, et al. Early life risk factors for obesity in childhood: cohort study. Br Med J 2005;330:1357.
[9] Taveras EM, Rifas-Shiman SL, Oken E, et al. Short sleep duration in infancy and risk of childhood overweight. Arch Pediatr Adolesc Med 2008;162:305–11.
[10] Tikotzky L, De Marcas G, Har-Toov J, et al. Sleep and physical growth in infants during the first 6 months. J Sleep Res 2010;19:103–10.
[11] Spiegel K, Tasali E, Penev P, et al. Brief communication: sleep curtailment in healthy young men is associated with decreased leptin levels, elevated ghrelin levels, and increased hunger and appetite. Ann Intern Med 2004;141:846–50.
[12] Sassin JF, Parker DC, Mace JW, et al. Human growth hormone release: relation to slow-wave sleep and sleep-waking cycles. Science 1969;165:513–15.
[13] Agras WS, Hammer LD, McNicholas F, et al. Risk factors for childhood overweight: a prospective study from birth to 9.5 years. J Pediatr 2004;145:20–5.
[14] Steptoe A, Peacey V, Wardle J. Sleep duration and health in young adults. Arch Intern Med 2006;166:1689–92.
[15] Olds T, Blunden S, Petkov J, et al. The relationships between sex, age, geography and time in bed in adolescents: a meta-analysis of data from 23 countries. Sleep Med Rev 2010;14:371–8.
[16] Mindell JA, Sadeh A, Wiegand B, et al. Cross-cultural differences in infant and toddler sleep. Sleep Med 2010;11:274–80.
[17] Aishworiya R, Chan P, Kiing J, et al. Sleep behaviour in a sample of preschool children in Singapore. Ann Acad Med Singapore 2012;41:99–104.
[18] Iglowstein I, Jenni OG, Molinari L, et al. Sleep duration from infancy to adolescence: reference values and generational trends. Pediatrics 2003; 111:302–7.
[19] Soh SE, Tint MT, Gluckman PD, et al. Cohort profile: growing up in singapore towards healthy outcomes (GUSTO) birth cohort study. Int J Epidemiol 2014;43:1401–9.
[20] Sadeh AA. Brief screening questionnaire for infant sleep problems validation and findings for an internet sample. Pediatrics 2004;113.
[21] The World Health Organization. Indicators for assessing infant and young child feeding practices: part 1. WHO Press; 2008.
[22] Bonuck KA, Freeman K, Henderson J. Growth and growth biomarker changes after adenotonsillectomy: systematic review and metaanalysis. Arch Dis Child 2009;94:83–91.
[23] Gulliford MC, Price CE, Rona RJ, et al. Sleep habits and height at ages 5 to 11. Arch Dis Child 1990;65:119–22.